

SEQUENCE LISTING

<110> Clausen, Henrik Bennett, Eric P.

<120> METHODS TO IDENTIFY AGENTS MODULATING FUNCTIONS OF POLYPEPTIDE GALNAC-TRANSFERASES, PHARMACEUTICAL COMPOSITIONS COMPRISING SUCH AGENTS AND THE USE OF SUCH AGENTS FOR PREPARING MEDICAMENTS

- <130> 04305/100H154-US2
- <150> US 60/425,204
- <151> 2002-11-08
- <150> PCT/DK03/00763
- <151> 2003-11-07
- <160> 127
- <170> PatentIn version 3.1
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- <213> Artificial Sequence
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- <223> synthetic peptide
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Ala Pro Pro Ala 20

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Ala Pro Gly Ser Thr Ala Pro Pro 20

- <210> 3
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- <212> PRT
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Lys Pro Phe Ser Trp Tyr Leu Glu Asp Ile Tyr Pro Asp Ser Gln Ile
Pro Arg His Tyr Phe Ser Leu Gly Glu Ile Arg Asn Val Glu Thr Asn
Gln Cys Leu Asp Asn Met Ala Arg Lys Glu Asn Glu Lys Val Gly Ile
Phe Asn Cys His Gly Met Gly Gly Asn Gln Val Phe Ser Tyr Thr Ala
65 Asn Lys Glu Ile Arg Thr Asp Asp Leu Cys Leu Asp Val Ser Lys Leu
Asn Gly Pro Val Thr Met Leu Lys Cys His His Leu Lys Gly Asn Gln
Leu Trp Glu Tyr Asp Pro Val Lys Leu Thr Leu Gln His Val Asn Ser
Asn Gln Cys Leu Asp Lys Ala Thr Glu Glu Asp Ser Gln Val Pro Ser

Ile Arg Asp Cys Asn Gly Ser Arg Ser Gln Gln Trp Leu Leu Arg Asn
Inches Arg Asp Cys Asn Gly Ser Arg Ser Gln Gln Trp Leu Leu Arg Asn

Val Thr Leu Pro Glu Ile Phe

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Tyr Gly Asn Ile Gln Ser Arg Leu Glu Leu Arg Lys Lys Leu Ser Cys $1 \hspace{1cm} 10 \hspace{1cm} 15$

Lys Pro Phe Lys Trp Tyr Leu Glu Asn Val Tyr Pro Glu Leu Arg Val 20 25 30

Pro Asp His Gln Asp Ile Ala Phe Gly Ala Leu Gln Gln Gly Thr Asn 35 40 45

Cys Leu Asp Thr Leu Gly His Phe Ala Asp Gly Val Val Gly Val Tyr

55

50

60

Glu Cys His Asn Ala Gly Gly Asn Gln Glu Trp Ala Leu Thr Lys Glu 65 70 75 80

Lys Ser Val Lys His Met Asp Leu Cys Leu Thr Val Val Asp Arg Ala 85 90 95

Pro Gly Ser Leu Ile Lys Leu Gln Gly Cys Arg Glu Asn Asp Ser Arg 100 105 110

Gln Lys Trp Glu Gln Ile Glu Gly Asn Ser Lys Leu Arg His Val Gly 115 120 125

Ser Asn Leu Cys Leu Asp Ser Arg Thr Ala Lys Ser Gly Gly Leu Ser 130 140

Val Glu Val Cys Gly Pro Ala Leu Ser Gln Gln Trp Lys Phe Thr Leu 145 150 155 160

Asn Leu Gln Gln

<210> 5

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<400> 5

Phe Gly Asp Leu Ser Lys Arg Phe Glu Ile Lys His Arg Leu Arg Cys 1 10 15

Lys Asn Phe Thr Trp Tyr Leu Asn Asn Ile Tyr Pro Glu Val Tyr Val 20 25 30

Pro Asp Leu Asn Pro Val Ile Ser Gly Tyr Ile Lys Ser Val Gly Gln 35 40 45

Pro Leu Cys Leu Asp Val Gly Glu Asn Asn Gln Gly Gly Lys Pro Leu 50 60

Ile Met Tyr Thr Cys His Gly Leu Gly Gly Asn Gln Tyr Phe Glu Tyr 65 70 75 80

Ser Ala Gln His Glu Ile Arg His Asn Ile Gln Lys Glu Leu Cys Leu 85 90 95

His Ala Ala Gln Gly Leu Val Gln Leu Lys Ala Cys Thr Tyr Lys Gly
100 105 110

His Lys Thr Val Val Thr Gly Glu Gln Ile Trp Glu Ile Gln Lys Asp 115 120 125

Gln Leu Leu Tyr Asn Pro Phe Leu Lys Met Cys Leu Ser Ala Asn Gly 130 135 140

Glu His Pro Ser Leu Val Ser Cys Asn Pro Ser Asp Pro Leu Gln Lys 145 150 155 160

Trp Ile Leu Ser Gln Asn Asp 165

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Ala Tyr Gly Asp Ile Ser Glu Arg Lys Leu Leu Arg Glu Arg Leu Arg
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Cys Lys Ser Phe Asp Trp Tyr Leu Lys Asn Val Phe Pro Asn Leu His 20 25 30

Val Pro Glu Asp Arg Pro Gly Trp His Gly Ala Ile Arg Ser Arg Gly 35 40 45

Ile Ser Ser Glu Cys Leu Asp Tyr Asn Ser Pro Asp Asn Asn Pro Thr 50 60

Gly Ala Asn Leu Ser Leu Phe Gly Cys His Gly Gln Gly Gly Asn Gln 65 70 75 80

Phe Phe Glu Tyr Thr Ser Asn Lys Glu Ile Arg Phe Asn Ser Val Thr 85 90 95

Glu Leu Cys Ala Glu Val Pro Glu Gln Lys Asn Tyr Val Gly Met Gln 100 105 110

Asn Cys Pro Lys Asp Gly Phe Pro Val Pro Ala Asn Ile Ile Trp His 115 120 125

Phe Lys Glu Asp Gly Thr Ile Phe His Pro His Ser Gly Leu Cys Leu 130 140

Ser Ala Tyr Arg Thr Pro Glu Gly Arg Pro Asp Val Gln Met Arg Thr 145 150 155 160 Cys Asp Ala Leu Asp Lys Asn Gln Ile Trp Ser Phe Glu Lys 165 170

<210> 7

<211> 168

<212> PRT

<213> Homo sapiens

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Asp Val Gly Asn Leu Thr Gln Gln Arg Glu Leu Arg Lys Lys Leu Lys 1 5 10 15

Cys Lys Ser Phe Lys Trp Tyr Leu Glu Asn Val Phe Pro Asp Leu Arg 20 25 30

Ala Pro Ile Val Arg Ala Ser Gly Val Leu Ile Asn Val Ala Leu Gly 35 40 45

Lys Cys Ile Ser Ile Glu Asn Thr Thr Val Ile Leu Glu Asp Cys Asp 50 55 60

Gly Ser Lys Glu Leu Gln Gln Phe Asn Tyr Thr Trp Leu Arg Leu Ile 65 70 75 80

Lys Cys Gly Glu Trp Cys Ile Ala Pro Ile Pro Asp Lys Gly Ala Val 85 90 95

Arg Leu His Pro Cys Asp Asn Arg Asn Lys Gly Leu Lys Trp Leu His
100 105 110

Lys Ser Thr Ser Val Phe His Pro Glu Leu Val Asn His Ile Val Phe 115 120 125

Glu Asn Asn Gln Gln Leu Leu Cys Leu Glu Gly Asn Phe Ser Gln Lys 130 135 140

Ile Leu Lys Val Ala Ala Cys Asp Pro Val Lys Pro Tyr Gln Lys Trp
145 150 155 160

Lys Phe Glu Lys Tyr Tyr Glu Ala 165

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<213> Homo sapiens

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Ser Phe Gly Asp Ile Ser Glu Arg Leu Gln Leu Arg Glu Gln Leu His

Cys His Asn Phe Ser Trp Tyr Leu His Asn Val Tyr Pro Glu Met Phe 20 25 30

Val Pro Asp Leu Thr Pro Thr Phe Tyr Gly Ala Ile Lys Asn Leu Gly 35 40 45

Thr Asn Gln Cys Leu Asp Val Gly Glu Asn Asn Arg Gly Gly Lys Pro 50 60

Leu Ile Met Tyr Ser Cys His Gly Leu Gly Gly Asn Gln Tyr Phe Glu 65 70 75 80

Tyr Thr Thr Gln Arg Asp Leu Arg His Asn Ile Ala Lys Gln Leu Cys 85 90 95

Leu His Val Ser Lys Gly Ala Leu Gly Leu Gly Ser Cys His Phe Thr 100 105 110

Gly Lys Asn Ser Gln Val Pro Lys Asp Glu Glu Trp Glu Leu Ala Gln 115 120 125

Asp Gln Leu Ile Arg Asn Ser Gly Ser Gly Thr Cys Leu Thr Ser Gln 130 140

Asp Lys Lys Pro Ala Met Ala Pro Cys Asn Pro Ser Asp Pro His Gln 145 150 155 160

Leu Trp Leu Phe Val

<210>

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<212> PRT

<213> Homo sapiens

<400> 9

Tyr Gly Asp Ile Ser Glu Leu Lys Lys Phe Arg Glu Asp His Asn Cys 1 10 15

Gln Ser Phe Lys Trp Phe Met Glu Glu Ile Ala Tyr Asp Ile Thr Ser 20 25 30

His Tyr Pro Leu Pro Pro Lys Asn Val Asp Trp Gly Glu Ile Arg Gly 35 40 45

Phe Glu Thr Ala Tyr Cys Ile Asp Ser Met Gly Lys Thr Asn Gly Gly 50 55 60

Phe Val Glu Leu Gly Pro Cys His Arg Met Gly Gly Asn Gln Leu Phe 65 70 75 80

Arg Ile Asn Glu Ala Asn Gln Leu Met Gln Tyr Asp Gln Cys Leu Thr 85 90 95

Lys Gly Ala Asp Gly Ser Lys Val Met Ile Thr His Cys Asn Leu Asn 100 105 110

Glu Phe Lys Glu Trp Gln Tyr Phe Lys Asn Leu His Arg Phe Thr His 125

Ile Pro Ser Gly Lys Cys Leu Asp Arg Ser Glu Val Leu His Gln Val 130 135 140

Phe Ile Ser Asn Cys Asp Ser Ser Lys Thr Thr Gln Lys Trp Glu Met 145 150 155 160

Asn Asn Ile His Ser Val 165

<210> 10

<211> 179 <212> PRT

<213> Homo sapiens

<400> 10

Phe Gly Asp Val Ser Ser Arg Met Ala Leu Arg Glu Lys Leu Lys Cys $1 \hspace{1cm} 10 \hspace{1cm} 15$

Lys Thr Phe Asp Trp Tyr Leu Lys Asn Val Tyr Pro Leu Lys Pro $20 \ 25 \ 30$

Leu His Thr Ile Val Gly Tyr Gly Arg Met Lys Asn Leu Leu Asp Glu 35 40 45

Asn Val Cys Leu Asp Gln Gly Pro Val Pro Gly Asn Thr Pro Ile Met 50 55 60

Tyr Tyr Cys His Glu Phe Ser Ser Gln Asn Val Tyr Tyr His Leu Thr 65 70 75 80

Gly Glu Leu Tyr Val Gly Gln Leu Ile Ala Glu Ala Ser Ala Ser Asp 85 90 95

Arg Cys Leu Thr Asp Pro Gly Lys Ala Glu Lys Pro Thr Leu Glu Pro 100 105 110 Cys Ser Lys Ala Ala Lys Asn Arg Leu His Ile Tyr Trp Asp Phe Lys 115 120 125

Pro Gly Gly Ala Val Ile Asn Arg Asp Thr Lys Arg Cys Leu Glu Met 130 140

Lys Lys Asp Leu Leu Gly Ser His Val Leu Val Leu Gln Thr Cys Ser 145 150 155 160

Thr Gln Val Trp Glu Ile Gln His Thr Val Arg Asp Trp Gly Gln Thr 165 170 175

Asn Ser Gln

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<212> PRT

<213> Homo sapiens

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Phe Gly Asp Val Ser Glu Arg Leu Ala Leu Arg Gln Arg Leu Lys Cys
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Arg Ser Phe Lys Trp Tyr Leu Glu Asn Val Tyr Pro Glu Met Arg Val 20 25 30

Tyr Asn Asn Thr Leu Thr Tyr Gly Glu Val Arg Asn Ser Lys Ala Ser 35 40 45

Ala Tyr Cys Leu Asp Gln Gly Ala Glu Asp Gly Asp Arg Ala Ile Leu 50 60

Tyr Pro Cys His Gly Met Ser Ser Gln Leu Val Arg Tyr Ser Ala Asp 65 70 75 80

Gly Leu Leu Gln Leu Gly Pro Leu Gly Ser Thr Ala Phe Leu Pro Asp 85 90 95

Ser Lys Cys Leu Val Asp Asp Gly Thr Gly Arg Met Pro Thr Leu Lys 100 105 110

Arg Cys Glu Asp Val Ala Arg Pro Thr Gln Arg Leu Trp Asp Phe Thr 115 120 125

Gln Ser Gly Pro Ile Val Ser Arg Ala Thr Gly Arg Cys Leu Glu Val 130 135 140

Glu Met Ser Lys Asp Ala Asn Phe Gly Leu Arg Leu Val Val Gln Arg 145 150 155 160

Cys Ser Gly Gln Lys Trp Met Ile Arg Asn Trp Ile Lys His Ala Arg 165 170 175

His

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PRT

<213> Homo sapiens

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Ala Gly Asp Val Ala Val Gln Lys Lys Leu Arg Ser Ser Leu Asn Cys
1 10 15

Lys Ser Phe Lys Trp Phe Met Thr Lys Ile Ala Trp Asp Leu Pro Lys 20 25 30

Phe Tyr Pro Pro Val Glu Pro Pro Ala Ala Ala Trp Gly Glu Ile Arg 35 40 45

Asn Val Gly Thr Gly Leu Cys Ala Asp Thr Lys His Gly Ala Leu Gly 50 60

Ser Pro Leu Arg Leu Glu Gly Cys Val Arg Gly Arg Gly Glu Ala Ala 65 70 75 80

Trp Asn Asn Met Gln Val Phe Thr Phe Thr Trp Arg Glu Asp Ile Arg 85 90 95

Pro Gly Asp Pro Gln His Thr Lys Lys Phe Cys Phe Asp Ala Ile Ser 100 105 110

His Thr Ser Pro Val Thr Leu Tyr Asp Cys His Ser Met Lys Gly Asn 115 120 125

Gln Leu Trp Lys Tyr Arg Lys Asp Lys Thr Leu Tyr His Pro Val Ser 130 135 140

Gly Ser Cys Met Asp Cys Ser Glu Ser Asp His Arg Ile Phe Met Asn 145 150 155

Thr Cys Asn Pro Ser Ser Leu Thr Gln Gln Trp Leu Phe Glu His Thr 165 170 175

Asn Ser Thr Val Leu Glu Lys Phe Asn Arg Asn

180 185

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<213> Homo sapiens

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Asn Ile Ser Glu Arg Val Glu Leu Arg Lys Leu Gly Cys Lys Ser 1 10 15

Phe Lys Trp Tyr Leu Asp Asn Val Tyr Pro Glu Met Gln Ile Ser Gly 20 25 30

Ser His Ala Lys Pro Gln Gln Pro Ile Phe Val Asn Arg Gly Pro Lys 35 40 45

Pro Lys Val Leu Gln Arg Gly Arg Leu Tyr His Leu Gln Thr Asn 50 60

Lys Cys Leu Val Ala Gln Gly Arg Pro Ser Gln Lys Gly Gly Leu Val 65 70 75 80

Val Leu Lys Ala Cys Asp Tyr Ser Asp Pro Asn Gln Ile Trp Ile Tyr 85 90 95

Asn Glu Glu His Glu Leu Val Leu Asn Ser Leu Leu Cys Leu Asp Met 100

Ser Glu Thr Arg Ser Ser Asp Pro Pro Arg Leu Met Lys Cys His Gly
115 120 125 115

Ser Gly Gly Ser Gln Gln Trp Thr Phe Gly Lys Asn Asn Arg Leu Tyr 130 140

Gln Val Ser Val Gly Gln Cys Leu Arg Ala Val Asp Pro Leu Gly Gln 145 150 155 160

Lys Gly Ser Val Ala Met Ala Ile Cys Asp Gly Ser Ser Ser Gln Gln 165 170 175

Trp His Leu Glu Gly 180

<400> 14

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<213> Homo sapiens

Asp Val Thr Glu Arg Lys Gln Leu Arg Asp Lys Leu Gln Cys Lys Asp
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Phe Lys Trp Phe Leu Glu Thr Val Tyr Pro Glu Leu His Val Pro Glu 20 25 30

Asp Arg Pro Gly Phe Phe Gly Met Leu Gln Asn Lys Gly Leu Thr Asp 35 40 45

Tyr Cys Phe Asp Tyr Asn Pro Pro Asp Glu Asn Gln Ile Val Gly His 50 60

Gln Val Ile Leu Tyr Leu Cys His Gly Met Gly Gln Asn Gln Phe Phe 65 70 75 80

Glu Tyr Thr Ser Gln Lys Glu Ile Arg Tyr Asn Thr His Gln Pro Glu 85 90 95

Gly Cys Ile Ala Val Glu Ala Gly Met Asp Thr Leu Ile Met His Leu 100 105 110

Cys Glu Glu Thr Ala Pro Glu Asn Gln Lys Phe Ile Leu Gln Glu Asp

Gly Ser Leu Phe His Glu Gln Ser Lys Lys Cys Val Gln Ala Ala Arg 130 135 140

Lys Glu Ser Ser Asp Ser Phe Val Pro Leu Leu Arg Asp Cys Thr Asn

Ser Asp His Gln Lys Trp Phe Phe Lys Glu Arg Met Leu 165 170

<400> 15

Glu Lys Pro Asp Cys Met Glu Arg Leu Gln Leu Gln Arg Arg Leu Gly
1 10 15

Cys Arg Thr Phe His Trp Phe Leu Ala Asn Val Tyr Pro Glu Leu Tyr 20 25 30

Pro Ser Glu Pro Arg Pro Ser Phe Ser Gly Lys Leu His Asn Thr Gly 35 40 45

<210>

¹⁵ 174 <211>

<212> **PRT**

<213> Homo sapiens

Leu Gly Leu Cys Ala Asp Cys Gln Ala Glu Gly Asp Ile Leu Gly Cys 50 60

Pro Met Val Leu Ala Pro Cys Ser Asp Ser Arg Gln Gln Gln Tyr Leu 65 70 75 80

Gln His Thr Ser Arg Lys Glu Ile His Phe Gly Ser Pro Gln His Leu 85 90 95

Cys Phe Ala Val Arg Gln Glu Gln Val Ile Leu Gln Asn Cys Thr Glu
100 105 110

Glu Gly Leu Ala Ile His Gln Gln His Trp Asp Phe Gln Glu Asn Gly 115 120 125

Met Ile Val His Ile Leu Ser Gly Lys Cys Met Glu Ala Val Val Gln 130 135 140

Glu Asn Asn Lys Asp Leu Tyr Leu Arg Pro Cys Asp Gly Lys Ala Arg 145 150 155 160

Gln Gln Trp Arg Phe Asp Gln Ile Asn Ala Val Asp Glu Arg

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Tyr Gly Asp Val Ser Val Arg Lys Thr Leu Arg Glu Asn Leu Lys Cys
1 10 15

Lys Pro Phe Ser Trp Tyr Leu Glu Asn Ile Tyr Pro Asp Ser Gln Ile 20 25 30

Pro Arg Arg Tyr Tyr Ser Leu Gly Glu Ile Arg Asn Val Glu Thr Asn 35 40 45

Gln Cys Leu Asp Asn Met Gly Arg Lys Glu Asn Glu Lys Val Gly Ile 50 60

Phe Asn Cys His Gly Met Gly Gly Asn Gln Val Phe Ser Tyr Thr Ala 65 70 75 80

Asp Lys Glu Ile Arg Thr Asp Asp Leu Cys Leu Asp Val Ser Arg Leu 85 90 95

Asn Gly Pro Val Ile Met Leu Lys Cys His His Met Arg Gly Asn Gln

¹⁶

<210> <211> 165

<212> PRT

<213> Homo sapiens

100 105 110

Leu Trp Glu Tyr Asp Ala Glu Arg Leu Thr Leu Arg His Val Asn Ser 115 120 125

Asn Gln Cys Leu Asp Glu Pro Ser Glu Glu Asp Lys Met Val Pro Thr 130 135 140

Met Gln Asp Cys Ser Gly Ser Arg Ser Gln Gln Trp Leu Leu Arg Asn 145 150 155 160

Met Thr Leu Gly Thr 165

<210> 17

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<212> PRT

<213> Homo sapiens

<400> 17

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Phe Gly Asn Val Glu Ser Arg Leu Asp Leu Arg Lys Asn Leu Arg Cys 1 10 15

Gln Ser Phe Lys Trp Tyr Leu Glu Asn Ile Tyr Pro Glu Leu Ser Ile 20 25 30

Pro Lys Glu Ser Ser Ile Gln Lys Gly Asn Ile Arg Gln Arg Gln Lys 35 40 45

Cys Leu Glu Ser Gln Arg Gln Asn Asn Gln Glu Thr Pro Asn Leu Lys 50 60

Leu Ser Pro Cys Ala Lys Val Lys Gly Glu Asp Ala Lys Ser Gln Val 65 70 75 80

Trp Ala Phe Thr Tyr Thr Gln Lys Ile Leu Gln Glu Glu Leu Cys Leu 85 90 95

Ser Val Ile Thr Leu Phe Pro Gly Ala Pro Val Val Leu Val Leu Cys 100 105 110

Lys Asn Gly Asp Asp Arg Gln Gln Trp Thr Lys Thr Gly Ser His Ile 115 120 125

Glu His Ile Ala Ser His Leu Cys Leu Asp Thr Asp Met Phe Gly Asp 130 135 140

Gly Thr Glu Asn Gly Lys Glu Ile Gly Val Asn Pro Cys Glu Ser Ser 145 150 155 160 Leu Met Ser Gln His Trp Asp Met Val Ser Ser 165 170

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<213> Homo sapiens

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Ser Val Ala Thr Arg Ile Glu Gln Arg Lys Lys Met Asn Cys Lys Ser $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$

Phe Arg Trp Tyr Leu Glu Asn Val Tyr Pro Glu Leu Thr Val Pro Val 20 25 30

Lys Glu Ala Leu Pro Gly Ile Ile Lys Gln Gly Val Asn Cys Leu Glu 35 40 45

Ser Gln Gly Gln Asn Thr Ala Gly Asp Phe Leu Leu Gly Met Gly Ile $50 \hspace{1cm} 55 \hspace{1cm} 60$

Cys Arg Gly Ser Ala Lys Asn Pro Gln Pro Ala Gln Ala Trp Leu Phe 65 70 75 80

Ser Asp His Leu Ile Gln Gln Gln Gly Lys Cys Leu Ala Ala Thr Ser 85 90 95

Thr Leu Met Ser Ser Pro Gly Ser Pro Val Ile Leu Gln Met Cys Asn $100 \hspace{1cm} 105 \hspace{1cm} 110$

Pro Arg Glu Gly Lys Gln Lys Trp Arg Arg Lys Gly Ser Phe Ile Gln 115 120 125

His Ser Val Ser Gly Leu Cys Leu Glu Thr Lys Pro Ala Gln Leu Val 130 135 140

Thr Ser Lys Cys Gln Ala Asp Ala Gln Ala Gln Gln Trp Gln Leu Leu 145 150 155 160

Pro His Thr

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cgtggg	gccg	gggccggggc	tgccgagccg	ggacccccgc	gcaccccgcg	ccccgggcgg	180
cgcgag	ccgg	tcatgccgcg	gccgccggtg	ccggcgaacg	cgctgggcgc	gcggggcgag	240
gcggtg	cggc	tgcagctgca	gggcgaggag	ctgcggctgc	aggaggagag	cgtgcggctg	300
caccaga	atta	acatctacct	cagcgaccgc	atctcactgc	accgccgcct	gcccgagcgc	360
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Met Trp Gly Arg Thr Ala Arg Arg Cys Pro Arg Glu Leu Arg Arg 1 5 10 15

Gly Arg Glu Ala Leu Leu Val Leu Leu Ala Leu Leu Ala Gly 20 25 30

Leu Gly Ser Val Leu Arg Ala Gln Arg Gly Ala Gly Ala Gly Ala Ala 35

Glu Pro Gly Pro Pro Arg Thr Pro Arg Pro Gly Arg Arg Glu Pro Val 50 60

Met Pro Arg Pro Pro Val Pro Ala Asn Ala Leu Gly Ala Arg Gly Glu 65 75 80 22

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⁵⁸¹

PRT

Homo sapiens

<400> 55

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Gly Glu Asn Leu Glu Phe Ser Phe Arg Ile Trp Gln Cys Gly Gly Val 340 345 350 Leu Glu Thr His Pro Cys Ser His Val Gly His Phe Ser Pro Ser Lys 355 360 365 Leu Pro Thr Pro Arg Asn Lys Ala Leu Ala Asn Ser Val Arg Ala Ala 370 380 Glu Val Trp Met Asp Glu Phe Lys Glu Leu Tyr Tyr His Arg Asn Pro 385 390 395 400 Arg Ala Arg Leu Glu Pro Phe Gly Asp Val Thr Glu Arg Lys Gln Leu Arg Asp Lys Leu Gln Cys Lys Asp Phe Lys Trp Phe Leu Glu Thr Val Tyr Pro Glu Leu His Val Pro Glu Asp Arg Pro Gly Phe Phe Gly Met Leu Gln Asn Lys Gly Leu Thr Asp Tyr Cys Phe Asp Tyr Asn Pro Pro Asp Glu Asn Gln Ile Val Gly His Gln Val Ile Leu Tyr Leu Cys His 465 470 475 480 Gly Met Gly Gln Asn Gln Phe Phe Glu Tyr Thr Ser Gln Lys Glu Ile Arg Tyr Asn Thr His Gln Pro Glu Gly Cys Ile Ala Val Glu Ala Gly 500 510 Met Asp Thr Leu Ile Met His Leu Cys Glu Glu Thr Ala Pro Glu Asn Gln Lys Phe Ile Leu Gln Glu Asp Gly Ser Leu Phe His Glu Gln Ser 530 540 Lys Lys Cys Val Gln Ala Ala Arg Lys Glu Ser Ser Asp Ser Phe Val 545 550 555 560 Pro Leu Leu Arg Asp Cys Thr Asn Ser Asp His Gln Lys Trp Phe Phe 565 570 575 Lys Glu Arg Met Leu 580

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<211> 639

<212> PRT <213> Homo sapiens

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Leu His Pro Pro His His Thr Leu His Gln Thr Val Thr Ala Gln Ala 35 40 45

Ser Lys His Ser Pro Glu Ala Arg Tyr Arg Leu Asp Phe Gly Glu Ser 50 55 60

Gln Asp Trp Val Leu Glu Ala Glu Asp Glu Gly Glu Glu Tyr Ser Pro 65 70 75 80

Leu Glu Gly Leu Pro Pro Phe Ile Ser Leu Arg Glu Asp Gln Leu Leu 85 90 95

Val Ala Val Ala Leu Pro Gln Ala Arg Arg Asn Gln Ser Gln Gly Arg 100 105 110

Arg Gly Gly Ser Tyr Arg Leu Ile Lys Gln Pro Arg Arg Gln Asp Lys
115 120 125

Glu Ala Pro Lys Arg Asp Trp Gly Ala Asp Glu Asp Gly Glu Val Ser 130 135 140

Glu Glu Glu Leu Thr Pro Phe Ser Leu Asp Pro Arg Gly Leu Gln 155 160

Glu Ala Leu Ser Ala Arg Ile Pro Leu Gln Arg Ala Leu Pro Glu Val 165 170 175

Arg His Pro Leu Cys Leu Gln Gln His Pro Gln Asp Ser Leu Pro Thr 180 185 190 Ala Ser Val Ile Leu Cys Phe His Asp Glu Ala Trp Ser Thr Leu Leu 195 200 205 Thr Val His Ser Ile Leu Asp Thr Val Pro Arg Ala Phe Leu Lys 210 220 Glu Ile Ile Leu Val Asp Asp Leu Ser Gln Gln Gly Gln Leu Lys Ser 225 230 235 240 Ala Leu Ser Glu Tyr Val Ala Arg Leu Glu Gly Val Lys Leu Leu Arg 245 250 255 Ser Asn Lys Arg Leu Gly Ala Ile Arg Ala Arg Met Leu Gly Ala Thr 260 265 270 Arg Ala Thr Gly Asp Val Leu Val Phe Met Asp Ala His Cys Glu Cys Pro Gly Trp Leu Glu Pro Leu Leu Ser Arg Ile Ala Gly Asp Arg 290 295 300 Ser Arg Val Val Ser Pro Val Ile Asp Val Ile Asp Trp Lys Thr Phe 305 310 315 Gln Tyr Tyr Pro Ser Lys Asp Leu Gln Arg Gly Val Leu Asp Trp Lys 325 330 335 Leu Asp Phe His Trp Glu Pro Leu Pro Glu His Val Arg Lys Ala Leu 340 345 350 Gln Ser Pro Ile Ser Pro Ile Arg Ser Pro Val Val Pro Gly Glu Val 355 360 365 Val Ala Met Asp Arg His Tyr Phe Gln Asn Thr Gly Ala Tyr Asp Ser 370 380 Leu Met Ser Leu Arg Gly Gly Glu Asn Leu Glu Leu Ser Phe Lys Ala 385 390 395 400 Trp Leu Cys Gly Gly Ser Val Glu Ile Leu Pro Cys Ser Arg Val Gly
405 410 415 His Ile Tyr Gln Asn Gln Asp Ser His Ser Pro Leu Asp Gln Glu Ala Thr Leu Arg Asn Arg Val Arg Ile Ala Glu Thr Trp Leu Gly Ser Phe 435 440 445

Lys Glu Thr Phe Tyr Lys His Ser Pro Glu Ala Phe Ser Leu Ser Lys Ala Glu Lys Pro Asp Cys Met Glu Arg Leu Gln Leu Gln Arg Arg Leu 465 470 475 480 Gly Cys Arg Thr Phe His Trp Phe Leu Ala Asn Val Tyr Pro Glu Leu Tyr Pro Ser Glu Pro Arg Pro Ser Phe Ser Gly Lys Leu His Asn Thr Gly Leu Gly Leu Cys Ala Asp Cys Gln Ala Glu Gly Asp Ile Leu Gly 515 520 525 Cys Pro Met Val Leu Ala Pro Cys Ser Asp Ser Arg Gln Gln Gln Tyr 530 540 Leu Gln His Thr Ser Arg Lys Glu Ile His Phe Gly Ser Pro Gln His 545 550 555 560 Leu Cys Phe Ala Val Arg Gln Glu Gln Val Ile Leu Gln Asn Cys Thr 565 570 575 Glu Glu Gly Leu Ala Ile His Gln Gln His Trp Asp Phe Gln Glu Asn 580 Gly Met Ile Val His Ile Leu Ser Gly Lys Cys Met Glu Ala Val Val Gln Glu Asn Asn Lys Asp Leu Tyr Leu Arg Pro Cys Asp Gly Lys Ala 610 615 620 Arg Gln Gln Trp Arg Phe Asp Gln Ile Asn Ala Val Asp Glu Arg 625 630 635

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<212> DNA

<213> Homo sapiens

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Trp Val Leu Val Asp Val Phe Leu Leu Leu Tyr Phe Ser Glu Cys Asn 20 25 30

Lys Cys Asp Asp Lys Lys Glu Arg Ser Leu Leu Pro Ala Leu Arg Ala 35 40 45

PRT Homo sapiens

<400> 59

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30

Ile Gly Thr Tyr Asp Ala Gly Met Asp Ile Trp Gly Gly Glu Asn Leu 305 310 315 320 Glu Met Ser Phe Arg Ile Trp Gln Cys Gly Gly Ser Leu Glu Ile Val 325 330 335 Thr Cys Ser His Val Gly His Val Phe Arg Lys Ala Thr Pro Tyr Thr Phe Pro Gly Gly Thr Gly His Val Ile Asn Lys Asn Asn Arg Arg Leu 355 360 365 Ala Glu Val Trp Met Asp Glu Phe Lys Asp Phe Phe Tyr Ile Ile Ser Pro Gly Val Val Lys Val Asp Tyr Gly Asp Val Ser Val Arg Lys Thr Leu Arg Glu Asn Leu Lys Cys Lys Pro Phe Ser Trp Tyr Leu Glu Asn Ile Tyr Pro Asp Ser Gln Ile Pro Arg Arg Tyr Tyr Ser Leu Gly Glu 420 430 Ile Arg Asn Val Glu Thr Asn Gln Cys Leu Asp Asn Met Gly Arg Lys 435 Glu Asn Glu Lys Val Gly Ile Phe Asn Cys His Gly Met Gly Gly Asn 450 460 Gln Val Phe Ser Tyr Thr Ala Asp Lys Glu Ile Arg Thr Asp Asp Leu 465 470 475 480 470 465 Cys Leu Asp Val Ser Arg Leu Asn Gly Pro Val Ile Met Leu Lys Cys His His Met Arg Gly Asn Gln Leu Trp Glu Tyr Asp Ala Glu Arg Leu 500 505 510 Thr Leu Arg His Val Asn Ser Asn Gln Cys Leu Asp Glu Pro Ser Glu 515 520 525 Glu Asp Lys Met Val Pro Thr Met Gln Asp Cys Ser Gly Ser Arg Ser 530 540 Gln Gln Trp Leu Leu Arg Asn Met Thr Leu Gly Thr 545 550 555

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<211> 1911
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<211> 552 <212> PRT

<213> Homo sapiens

<400> 61

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Trp Asp Asp Leu Trp Asp Gln Phe Asp Glu Arg Arg Tyr Leu Asn Ala 50 60

Lys Lys Trp Arg Val Gly Asp Asp Pro Tyr Lys Leu Tyr Ala Phe Asn 65 70 75 80

Gln Arg Glu Ser Glu Arg Ile Ser Ser Asn Arg Ala Ile Pro Asp Thr 85 90 95

Arg His Leu Arg Cys Thr Leu Leu Val Tyr Cys Thr Asp Leu Pro Pro 100 105 110

Thr Ser Ile Ile Ile Thr Phe His Asn Glu Ala Arg Ser Thr Leu Leu 115 120 125

Arg Thr Ile Arg Ser Val Leu Asn Arg Thr Pro Thr His Leu Ile Arg 130 135 140

Glu Ile Ile Leu Val Asp Asp Phe Ser Asn Asp Pro Asp Asp Cys Lys 150 155 160

Gln Leu Ile Lys Leu Pro Lys Val Lys Cys Leu Arg Asn Asn Glu Arg 165 170 175

Gln Gly Leu Val Arg Ser Arg Ile Arg Gly Ala Asp Ile Ala Gln Gly 180 185 190

Thr Thr Leu Thr Phe Leu Asp Ser His Cys Glu Val Asn Arg Asp Trp Leu Gln Pro Leu Leu His Arg Val Lys Glu Asp Tyr Thr Arg Val Val 210 220 Cys Pro Val Ile Asp Ile Ile Asn Leu Asp Thr Phe Thr Tyr Ile Glu 225 230 235 240 Ser Ala Ser Glu Leu Arg Gly Gly Phe Asp Trp Ser Leu His Phe Gln 245 250 255 Trp Glu Gln Leu Ser Pro Glu Gln Lys Leu Gly Ala Trp Thr Pro Arg 260 265 270 Lys Pro Ile Arg Thr Pro Ile Ile Ala Gly Gly Leu Phe Val Ile Asp 275 280 285 Ala Trp Phe Asp Tyr Leu Gly Lys Tyr Asp Met Asp Met Asp Ile 290 295 300 Trp Gly Gly Glu Asn Phe Glu Ile Ser Phe Arg Val Trp Met Cys Gly 315 310 320 Gly Ser Leu Glu Ile Val Pro Cys Ser Arg Val Gly His Val Phe Arg 325 330 335 Lys Lys His Pro Tyr Val Phe Pro Asp Gly Asn Ala Asn Thr Tyr Ile 340 345 350 Lys Asn Thr Lys Arg Thr Ala Glu Val Trp Met Asp Glu Tyr Lys Gln Tyr Tyr Ala Ala Arg Pro Phe Ala Leu Glu Arg Pro Phe Gly Asn 370 375 380 Val Glu Ser Arg Leu Asp Leu Arg Lys Asn Leu Arg Cys Gln Ser Phe 385 395 400 Lys Trp Tyr Leu Glu Asn Ile Tyr Pro Glu Leu Ser Ile Pro Lys Glu 405 410 415 Ser Ser Ile Gln Lys Gly Asn Ile Arg Gln Arg Gln Lys Cys Leu Glu 420 425 430 Ser Gln Arg Gln Asn Asn Gln Glu Thr Pro Asn Leu Lys Leu Ser Pro 435 440 445 Cys Ala Lys Val Lys Gly Glu Asp Ala Lys Ser Gln Val Trp Ala Phe 450 455 460

Thr Tyr Thr Gln Gln Ile Leu Gln Glu Glu Leu Cys Leu Ser Val Ile 465 470 475 480

Thr Leu Phe Pro Gly Ala Pro Val Val Leu Val Leu Cys Lys Asn Gly 485 490 495

Asp Asp Arg Gln Gln Trp Thr Lys Thr Gly Ser His Ile Glu His Ile 500 505 510

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Gln Leu Arg Glu Asp Arg Thr Ile Pro Leu Ile Val Thr Gly Thr Pro 50 60

Ser Lys Gly Phe Asp Glu Lys Ala Tyr Leu Ser Ala Lys Gln Leu Lys 65 70 75 80

Ala Gly Glu Asp Pro Tyr Arg Gln His Ala Phe Asn Gln Leu Glu Ser 85 90 95

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Ile Glu Gln Arg Lys Lys Met Asn Cys Lys Ser Phe Arg Trp Tyr Leu 405 410 415

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Thr Ala Gly Asp Phe Leu Leu Gly Met Gly Ile Cys Arg Gly Ser Ala 450 455 460

Lys Asn Pro Gln Pro Ala Gln Ala Trp Leu Phe Ser Asp His Leu Ile 465 470 475 480

Gln Gln Gln Gly Lys Cys Leu Ala Ala Thr Ser Thr Leu Met Ser Ser 485 490 495

Pro Gly Ser Pro Val Ile Leu Gln Met Cys Asn Pro Arg Glu Gly Lys 500 510

Gln Lys Trp Arg Arg Lys Gly Ser Phe Ile Gln His Ser Val Ser Gly 515 520 525

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<213> Homo sapiens

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Pro Arg His Tyr Phe Ser Leu Gly Glu Ile Arg Asn Val Glu Thr Asn 35 40 45

Gln Cys Leu Asp Asn Met Ala Arg Lys Glu Asn Glu Lys Val Gly Ile 50 60

Phe Asn Cys His Gly Met Gly Gly Asn Gln Val Phe Ser Tyr Thr Ala 65 70 75 80

Asn Lys Glu Ile Arg Thr Asp Asp Leu Cys Leu Asp Val Ser Lys Leu 85 90 95

Asn Gly Pro Val Thr Met Leu Lys Cys His His Leu Lys Gly Asn Gln
100 105 110

Leu Trp Glu Tyr Asp Pro Val Lys Leu Thr Leu Gln His Val Asn Ser 115 120 125

Asn Gln Cys Leu Asp Lys Ala Thr Glu Glu Asp Ser Gln Val Pro Ser

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360	aagcgtggag	gcgggggcct	acggccaaga	ggacagtcgc	acctgtgcct	gtgggcagca
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Leu Gln Gln Gly Thr Asn Cys Leu Asp Thr Leu Gly His Phe Ala Asp 20 25 30

Gly Val Val Gly Val Tyr Glu Cys His Asn Ala Gly Gly Asn Gln Glu 35 40 45

Trp Ala Leu Thr Lys Glu Lys Ser Val Lys His Met Asp Leu Cys Leu 50 60

Thr Val Val Asp Arg Ala Pro Gly Ser Leu Ile Lys Leu Gln Gly Cys 65 70 75 80

Arg Glu Asn Asp Ser Arg Gln Lys Trp Glu Gln Ile Glu Gly Asn Ser 85 90 95

Lys Leu Arg His Val Gly Ser Asn Leu Cys Leu Asp Ser Arg Thr Ala 100 105 110

Lys Ser Gly Gly Leu Ser Val Glu Val Cys Gly Pro Ala Leu Ser Gln
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Gln Trp Lys Phe Thr Leu Asn Leu Gln Gln 130 135

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Pro Asp Leu Asn Pro Val Ile Ser Gly Tyr Ile Lys Ser Val Gly Gln 35 40 45

Pro Leu Cys Leu Asp Val Gly Glu Asn Asn Gln Gly Gly Lys Pro Leu 50 55 60

Ile Met Tyr Thr Cys His Gly Leu Gly Gly Asn Gln Tyr Phe Glu Tyr 65 70 75 80

Ser Ala Gln His Glu Ile Arg His Asn Ile Gln Lys Glu Leu Cys Leu 85 90 95

His Ala Ala Gln Gly Leu Val Gln Leu Lys Ala Cys Thr Tyr Lys Gly
100 105 110

His Lys Thr Val Val Thr Gly Glu Gln Ile Trp Glu Ile Gln Lys Asp 115 120 125

Gln Leu Leu Tyr Asn Pro Phe Leu Lys Met Cys Leu Ser Ala Asn Gly

Glu His Pro Ser Leu Val Ser Cys Asn Pro Ser Asp Pro Leu Gln Lys 145 150 155 160

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Cys Lys Ser Phe Asp Trp Tyr Leu Lys Asn Val Phe Pro Asn Leu His 20 25 30
Val Pro Glu Asp Arg Pro Gly Trp His Gly Ala Ile Arg Ser Arg Gly 35 40 45
Ile Ser Ser Glu Cys Leu Asp Tyr Asn Ser Pro Asp Asn Asn Pro Thr 50 60
Gly Ala Asn Leu Ser Leu Phe Gly Cys His Gly Gln Gly Gly Asn Gln 65 70 75 80
Phe Phe Glu Tyr Thr Ser Asn Lys Glu Ile Arg Phe Asn Ser Val Thr 85 90 95
Glu Leu Cys Ala Glu Val Pro Glu Gln Lys Asn Tyr Val Gly Met Gln 100 105 110

Asn Cys Pro Lys Asp Gly Phe Pro Val Pro Ala Asn Ile Ile Trp His Phe Lys Glu Asp Gly Thr Ile Phe His Pro His Ser Gly Leu Cys Leu Ser Ala Tyr Arg Thr Pro Glu Gly Arg Pro Asp Val Gln Met Arg Thr 150

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Ala Pro Ile Val Arg Ala Ser Gly Val Leu Ile Asn Val Ala Leu Gly 35 40 45

Lys Cys Ile Ser Ile Glu Asn Thr Thr Val Ile Leu Glu Asp Cys Asp 50 55 60

Gly Ser Lys Glu Leu Gln Gln Phe Asn Tyr Thr Trp Leu Arg Leu Ile 65 70 75 80 Lys Cys Gly Glu Trp Cys Ile Ala Pro Ile Pro Asp Lys Gly Ala Val 85 90 95 Arg Leu His Pro Cys Asp Asn Arg Asn Lys Gly Leu Lys Trp Leu His Lys Ser Thr Ser Val Phe His Pro Glu Leu Val Asn His Ile Val Phe Glu Asn Asn Gln Gln Leu Leu Cys Leu Glu Gly Asn Phe Ser Gln Lys 130 135 140 Ile Leu Lys Val Ala Ala Cys Asp Pro Val Lys Pro Tyr Gln Lys Trp 145 150 155 160 Lys Phe Glu Lys Tyr Tyr Glu Ala 165 <210> 106 <211> 498 DNA <212> <213> Homo sapiens <400> 106 60 tccttcggtg acatttcgga acgactgcag ctgagggaac aactgcactg tcacaacttt 120 180 tatggtgcca tcaagaacct cggcaccaac caatgcctgg atgtgggtga gaacaaccgc ggggggaagc ccctcatcat gtactcctgc cacggccttg gcggcaacca gtactttgag 240 300 tacacaactc agagggacct tcgccacaac atcgcaaagc agctgtgtct acatgtcagc 360 aagggtgctc tgggccttgg gagctgtcac ttcactggca agaatagcca ggtccccaag gacgaggaat gggaattggc ccaggatcag ctcatcagga actcaggatc tggtacctgc 420 480 ctgacatccc aggacaaaaa gccagccatg gccccctgca atcccagtga cccccatcag 498 ttgtggctct ttgtctag <210> 107 165 <212> PRT <213> Homo sapiens

Ser Phe Gly Asp Ile Ser Glu Arg Leu Gln Leu Arg Glu Gln Leu His 1 15

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Cys His Asn Phe Ser Trp Tyr Leu His Asn Val Tyr Pro Glu Met Phe 20 25 30 Val Pro Asp Leu Thr Pro Thr Phe Tyr Gly Ala Ile Lys Asn Leu Gly 35 40 45 Thr Asn Gln Cys Leu Asp Val Gly Glu Asn Asn Arg Gly Gly Lys Pro 50 60 Leu Ile Met Tyr Ser Cys His Gly Leu Gly Gly Asn Gln Tyr Phe Glu 65 70 75 80 Tyr Thr Thr Gln Arg Asp Leu Arg His Asn Ile Ala Lys Gln Leu Cys 85 90 95 Leu His Val Ser Lys Gly Ala Leu Gly Leu Gly Ser Cys His Phe Thr 100 105 110 Gly Lys Asn Ser Gln Val Pro Lys Asp Glu Glu Trp Glu Leu Ala Gln 115 120 125 Gln Leu Ile Arg Asn Ser Gly Ser Gly Thr Cys Leu Thr Ser Gln 130 140 Asp Lys Lys Pro Ala Met Ala Pro Cys Asn Pro Ser Asp Pro His Gln 145 150 155 160 Leu Trp Leu Phe Val <210> 108 501 DNA Homo sapiens <400> 108

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<211> 166

<212> PRT

<213> Homo sapiens

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His Tyr Pro Leu Pro Pro Lys Asn Val Asp Trp Gly Glu Ile Arg Gly 35 40 45

Phe Glu Thr Ala Tyr Cys Ile Asp Ser Met Gly Lys Thr Asn Gly Gly 50 55 60

Phe Val Glu Leu Gly Pro Cys His Arg Met Gly Gly Asn Gln Leu Phe 65 70 75 80

Arg Ile Asn Glu Ala Asn Gln Leu Met Gln Tyr Asp Gln Cys Leu Thr 85 90 95

Lys Gly Ala Asp Gly Ser Lys Val Met Ile Thr His Cys Asn Leu Asn $100 \hspace{1cm} 105 \hspace{1cm} 110$

Glu Phe Lys Glu Trp Gln Tyr Phe Lys Asn Leu His Arg Phe Thr His 115 120 125

Phe Ile Ser Asn Cys Asp Ser Ser Lys Thr Thr Gln Lys Trp Glu Met 145 150 155 160

Asn Asn Ile His Ser Val

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<212> DNA

<213> Homo sapiens

<400> 110

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atcatgtatt actgccatga attcagctca cagaatgtct actatcacct aactggggag 240 ctctatgtgg gacaactgat tgcagaggcc agtgctagtg atcgctgcct gacagaccct 300 ggcaaggcgg agaagcccac cttagaacca tgctccaagg cagctaagaa tagactgcat 360 atatattggg attttaaacc gggaggagct gtcataaaca gagataccaa gcggtgtctg 420 gagatgaaga aggatctttt gggtagccac gtgcttgtgc tccagacctg tagcacgcaa 480 gtgtgggaaa tccagcacac tgtcagagac tggggtcaga ccaacagcca gtga 534

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<400> 111

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Lys Thr Phe Asp Trp Tyr Leu Lys Asn Val Tyr Pro Leu Leu Lys Pro 20 25 30

Leu His Thr Ile Val Gly Tyr Gly Arg Met Lys Asn Leu Leu Asp Glu 35 40 45

Asn Val Cys Leu Asp Gln Gly Pro Val Pro Gly Asn Thr Pro Ile Met 50 55 60

Tyr Tyr Cys His Glu Phe Ser Ser Gln Asn Val Tyr Tyr His Leu Thr 65 70 75 80

Gly Glu Leu Tyr Val Gly Gln Leu Ile Ala Glu Ala Ser Ala Ser Asp 85 90 95

Arg Cys Leu Thr Asp Pro Gly Lys Ala Glu Lys Pro Thr Leu Glu Pro 100 105 110

Cys Ser Lys Ala Ala Lys Asn Arg Leu His Ile Tyr Trp Asp Phe Lys 115 120 125

Pro Gly Gly Ala Val Ile Asn Arg Asp Thr Lys Arg Cys Leu Glu Met 130 140

Lys Lys Asp Leu Leu Gly Ser His Val Leu Val Leu Gln Thr Cys Ser 145 150 155 160

Thr Gln Val Trp Glu Ile Gln His Thr Val Arg Asp Trp Gly Gln Thr 165 170 175 112 534

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Arg Ser Phe Lys Trp Tyr Leu Glu Asn Val Tyr Pro Glu Met Arg Val 20 25 30
Tyr Asn Asn Thr Leu Thr Tyr Gly Glu Val Arg Asn Ser Lys Ala Ser 35 40 45
Ala Tyr Cys Leu Asp Gln Gly Ala Glu Asp Gly Asp Arg Ala Ile Leu 50 60
Tyr Pro Cys His Gly Met Ser Ser Gln Leu Val Arg Tyr Ser Ala Asp 65 70 75 80
Gly Leu Leu Gln Leu Gly Pro Leu Gly Ser Thr Ala Phe Leu Pro Asp
85 90 95
Ser Lys Cys Leu Val Asp Asp Gly Thr Gly Arg Met Pro Thr Leu Lys
100 105 110
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60

120 180

240

300

360

420

480

Arg Cys Glu Asp Val Ala Arg Pro Thr Gln Arg Leu Trp Asp Phe Thr 115 120 125

Gln Ser Gly Pro Ile Val Ser Arg Ala Thr Gly Arg Cys Leu Glu Val 130 135 140

Glu Met Ser Lys Asp Ala Asn Phe Gly Leu Arg Leu Val Val Gln Arg 145 150 155 160

Cys Ser Gly Gln Lys Trp Met Ile Arg Asn Trp Ile Lys His Ala Arg 165 170 175

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<210> 115

<211> 187

<212> PRT

<213> Homo sapiens

<400> 115

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Lys Ser Phe Lys Trp Phe Met Thr Lys Ile Ala Trp Asp Leu Pro Lys 20 25 30

Phe Tyr Pro Pro Val Glu Pro Pro Ala Ala Ala Trp Gly Glu Ile Arg 35 40 45 Asn Val Gly Thr Gly Leu Cys Ala Asp Thr Lys His Gly Ala Leu Gly 50 60

Ser Pro Leu Arg Leu Glu Gly Cys Val Arg Gly Arg Gly Glu Ala Ala 65 70 75 80

Trp Asn Asn Met Gln Val Phe Thr Phe Thr Trp Arg Glu Asp Ile Arg 85 90 95

Pro Gly Asp Pro Gln His Thr Lys Lys Phe Cys Phe Asp Ala Ile Ser 100 105 110

His Thr Ser Pro Val Thr Leu Tyr Asp Cys His Ser Met Lys Gly Asn 115 120 125

Gln Leu Trp Lys Tyr Arg Lys Asp Lys Thr Leu Tyr His Pro Val Ser 130 135 140

Gly Ser Cys Met Asp Cys Ser Glu Ser Asp His Arg Ile Phe Met Asn 145 150 155 160

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Asn Ser Thr Val Leu Glu Lys Phe Asn Arg Asn

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549

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Homo sapiens

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<211> 181

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Phe Lys Trp Tyr Leu Asp Asn Val Tyr Pro Glu Met Gln Ile Ser Gly 20 25 30

Ser His Ala Lys Pro Gln Gln Pro Ile Phe Val Asn Arg Gly Pro Lys 35 40 45

Arg Pro Lys Val Leu Gln Arg Gly Arg Leu Tyr His Leu Gln Thr Asn 50 60

Lys Cys Leu Val Ala Gln Gly Arg Pro Ser Gln Lys Gly Gly Leu Val 65 70 75 80

Val Leu Lys Ala Cys Asp Tyr Ser Asp Pro Asn Gln Ile Trp Ile Tyr 85 90 95

Asn Glu Glu His Glu Leu Val Leu Asn Ser Leu Leu Cys Leu Asp Met 100 105 110

Ser Glu Thr Arg Ser Ser Asp Pro Pro Arg Leu Met Lys Cys His Gly 115 120 125

Ser Gly Gly Ser Gln Gln Trp Thr Phe Gly Lys Asn Asn Arg Leu Tyr 130 135 140

Gln Val Ser Val Gly Gln Cys Leu Arg Ala Val Asp Pro Leu Gly Gln 145 150 155 160

Lys Gly Ser Val Ala Met Ala Ile Cys Asp Gly Ser Ser Ser Gln Gln 165 170 175

Trp His Leu Glu Gly 180

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<211> 525

<212> DNA <213> Homo sapiens

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<400> 119

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Phe Lys Trp Phe Leu Glu Thr Val Tyr Pro Glu Leu His Val Pro Glu 20 25 30

Asp Arg Pro Gly Phe Phe Gly Met Leu Gln Asn Lys Gly Leu Thr Asp 35 40 45

Tyr Cys Phe Asp Tyr Asn Pro Pro Asp Glu Asn Gln Ile Val Gly His 50 55 60

Gln Val Ile Leu Tyr Leu Cys His Gly Met Gly Gln Asn Gln Phe Phe 65 70 75 80

Glu Tyr Thr Ser Gln Lys Glu Ile Arg Tyr Asn Thr His Gln Pro Glu 85 90 95

Gly Cys Ile Ala Val Glu Ala Gly Met Asp Thr Leu Ile Met His Leu 100 105 110

Cys Glu Glu Thr Ala Pro Glu Asn Gln Lys Phe Ile Leu Gln Glu Asp 115 120 125

Gly Ser Leu Phe His Glu Gln Ser Lys Lys Cys Val Gln Ala Ala Arg 130 135 140

Lys Glu Ser Ser Asp Ser Phe Val Pro Leu Leu Arg Asp Cys Thr Asn 145 150 155 160

<210> 119

<211> 173

<212> PRT

<213> Homo sapiens

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Cys Arg Thr Phe His Trp Phe Leu Ala Asn Val Tyr Pro Glu Leu Tyr 20 25 30
Pro Ser Glu Pro Arg Pro Ser Phe Ser Gly Lys Leu His Asn Thr Gly 35 40 45
Leu Gly Leu Cys Ala Asp Cys Gln Ala Glu Gly Asp Ile Leu Gly Cys 50 60
Pro Met Val Leu Ala Pro Cys Ser Asp Ser Arg Gln Gln Gln Tyr Leu 65 70 75 80
Gln His Thr Ser Arg Lys Glu Ile His Phe Gly Ser Pro Gln His Leu 85 90 95
Cys Phe Ala Val Arg Gln Glu Gln Val Ile Leu Gln Asn Cys Thr Glu 100 105 110

Glu Gly Leu Ala Ile His Gln Gln His Trp Asp Phe Gln Glu Asn Gly 115 120 125

Met Ile Val His Ile Leu Ser Gly Lys Cys Met Glu Ala Val Val Gln 130 135 140

Glu Asn Asn Lys Asp Leu Tyr Leu Arg Pro Cys Asp Gly Lys Ala Arg 145 150 155 160

Gln Gln Trp Arg Phe Asp Gln Ile Asn Ala Val Asp Glu Arg 165 170

<210> 122

<211> 498

<212> DNA

<213> Homo sapiens

<400> 122

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1 10 15

Lys Pro Phe Ser Trp Tyr Leu Glu Asn Ile Tyr Pro Asp Ser Gln Ile 20 25 30

Pro Arg Arg Tyr Tyr Ser Leu Gly Glu Ile Arg Asn Val Glu Thr Asn 35 40 45

Gln Cys Leu Asp Asn Met Gly Arg Lys Glu Asn Glu Lys Val Gly Ile 50 60

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<213> Homo sapiens

Phe Asn Cys His Gly Met Gly Gly Asn Gln Val Phe Ser Tyr Thr Ala 65 70 75 80 Asp Lys Glu Ile Arg Thr Asp Asp Leu Cys Leu Asp Val Ser Arg Leu 85 90 95 Asn Gly Pro Val Ile Met Leu Lys Cys His His Met Arg Gly Asn Gln 100 105 110 Leu Trp Glu Tyr Asp Ala Glu Arg Leu Thr Leu Arg His Val Asn Ser 115 120 125 Asn Gln Cys Leu Asp Glu Pro Ser Glu Glu Asp Lys Met Val Pro Thr Met Gln Asp Cys Ser Gly Ser Arg Ser Gln Gln Trp Leu Leu Arg Asn 150 155 160Met Thr Leu Gly Thr <210> 124 516 <211> <212> DNA <213> Homo sapiens <400> 124 60 tcgggaatgt tgagagcaga ttggacctga ggaagaatct gcgctgccag agcttcaagt 120 ggtacctgga gaatatctac cctgaactca gcatccccaa ggagtcctcc atccagaagg 180 gcaatatccg acagagacag aagtgcctgg aatctcaaag gcagaacaac caagaaaccc 240 caaacctaaa gttgagcccc tgtgccaagg tcaaaggcga agatgcaaag tcccaggtat 300 gggccttcac atacacccag aagatcctcc aggaggagct gtgcctgtca gtcatcacct tgttccctgg cgccccagtg gttcttgtcc tttgcaagaa tggagatgac cgacagcaat 360 ggaccaaaac tggttcccac atcgagcaca tagcatccca cctctgcctc gatacagata 420 480 tgttcggtga tggcaccgag aacggcaagg aaatcggcgt caacccatgt gagtcctcac tcatgagcca gcactgggac atggtgagtt cttgag 516 <210> 125 171 **PRT** Homo sapiens <400> 125 Phe Gly Asn Val Glu Ser Arg Leu Asp Leu Arg Lys Asn Leu Arg Cys
10 15

Gln	Ser	Phe	Lys 20	Trp	Tyr	Leu	Glu	Asn 25	Ile	Tyr	Pro	Glu	Leu 30	Ser	Ile		
Pro	Lys	Glu 35	Ser	Ser	Ile	Gln	Lys 40	Gly	Asn	Ile	Arg	G]n 45	Arg	Gln	Lys		
Cys	Leu 50	Glu	Ser	Gln	Arg	G]n 55	Asn	Asn	Gln	Glu	Thr 60	Pro	Asn	Leu	Lys		
Leu 65	Ser	Pro	Cys	Ala	Lys 70	∨al	Lys	Gly	Glu	Asp 75	Ala	Lys	Ser	Gln	Val 80		
Trp	Ala	Phe	Thr	Tyr 85	Thr	Gln	Lys	Ile	Leu 90	Gln	Glu	Glu	Leu	Cys 95	Leu		
Ser	٧a٦	Ile	Thr 100	Leu	Phe	Pro	Gly	Ala 105	Pro	val	val	Leu	val 110	Leu	Cys		
Lys	Asn	Gly 115	Asp	Asp	Arg	Gln	Gln 120	Trp	Thr	Lys	Thr	Gly 125	Ser	His	Il <u>'</u> e		
Glu	ніs 130	Ile	Ala	Ser	His	Leu 135	Cys	Leu	Asp	Thr	Asp 140	Met	Phe	Gly	Asp		
Gly 145	Thr	Glu	Asn	Gly	Lys 150	Glu	Ile	Gly	val	Asn 155	Pro	Cys	Glu	Ser	Ser 160		
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ggaa	atgg	gga 1	tctg	caga	gg g1	tctg	ccaa	g aad	cccg	cagc	ccg	cca	ggc .	atgg	tgttc	2	240
agt	gacc	acc 1	tcate	ccago	ca go	cagg	ggaa	g tg	cctg	gctg	cca	cctc	cac	cttaa	atgtcc	3	800
tcc	ctg	gat (ccca	agtca	at a	tgca	agat	g tg	caac	ccta	gaga	aaggo	caa	gcaga	aatgg	3	60
agga	agaa	aag g	gatci	tttca	at co	cagca	attca	a gto	cagt	ggcc	tct	gcct	gga	gacaa	agcct	4	120

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480

<210> 127

<211> 163

<212> PRT

<213> Homo sapiens

<400> 127

Ser Val Ala Thr Arg Ile Glu Gln Arg Lys Lys Met Asn Cys Lys Ser 1 10 15

Phe Arg Trp Tyr Leu Glu Asn Val Tyr Pro Glu Leu Thr Val Pro Val
20 25 30

Lys Glu Ala Leu Pro Gly Ile Ile Lys Gln Gly Val Asn Cys Leu Glu 35 40 45

Ser Gln Gly Gln Asn Thr Ala Gly Asp Phe Leu Leu Gly Met Gly Ile 50 60

Cys Arg Gly Ser Ala Lys Asn Pro Gln Pro Ala Gln Ala Trp Leu Phe 65 70 75 80

Ser Asp His Leu Ile Gln Gln Gln Gly Lys Cys Leu Ala Ala Thr Ser 85 90 95

Thr Leu Met Ser Ser Pro Gly Ser Pro Val Ile Leu Gln Met Cys Asn 100 105 110

Pro Arg Glu Gly Lys Gln Lys Trp Arg Arg Lys Gly Ser Phe Ile Gln 115 120 125

His Ser Val Ser Gly Leu Cys Leu Glu Thr Lys Pro Ala Gln Leu Val 130 135 140

Thr Ser Lys Cys Gln Ala Asp Ala Gln Ala Gln Gln Trp Gln Leu Leu 145 150 155 160

Pro His Thr